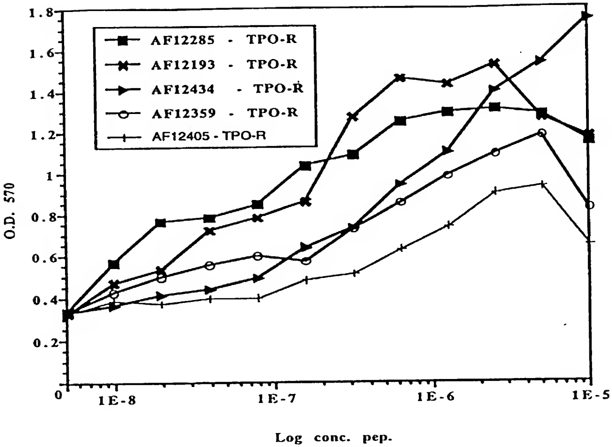


FIG. 1A

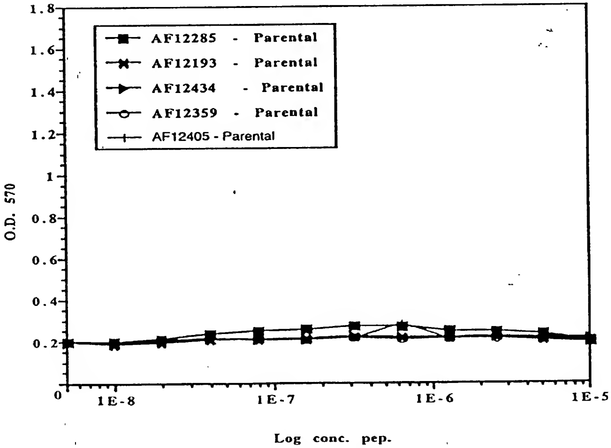


FIG. 1B

Express Mail Label No.
EV022025865US

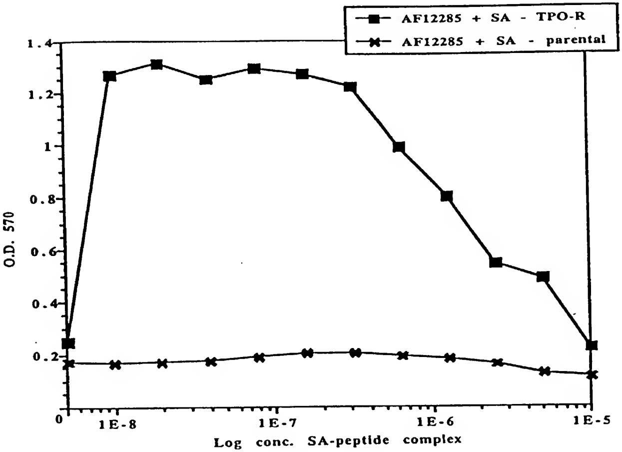

AF12285 + SA - TPO-R
AF12285 + SA - parental
1.4
1.2
1
0.8
0.6
0.4
0.2
0
O.D. 570
1E-8
1E-7
1E-6
1E-5
Log conc. SA-peptide complex

FIG. 2A

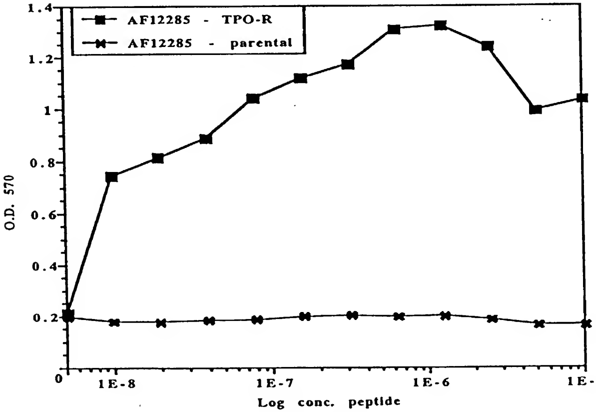

AF12285 - TPO-R
AF12285 - parental
1.4
1.2
1
0.8
0.6
0.4
0.2
0
O.D. 570
1E-8
1E-7
1E-6
1E-5
Log conc. peptide

FIG. 2B

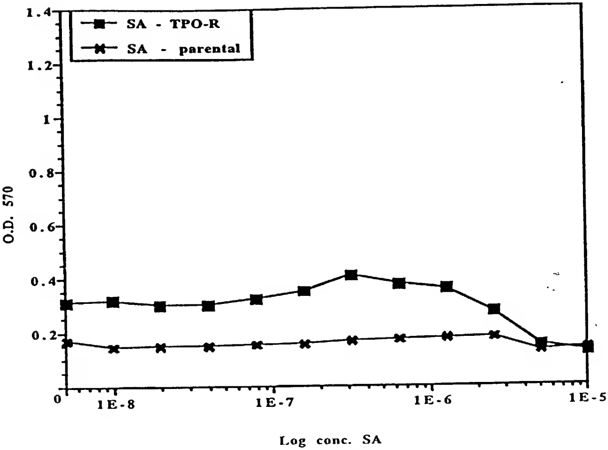

SA - TPO-R
SA - parental
O.D. 570
1.4
1.2
1
0.8
0.6
0.4
0.2
0
1E-8
1E-7
1E-6
1E-5
Log conc. SA

FIG. 2C

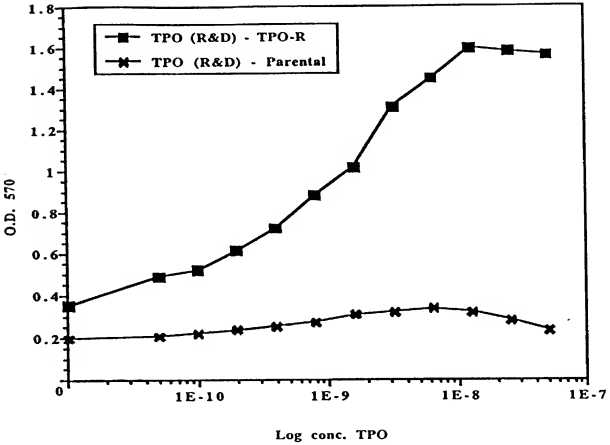

TPO (R&D) - TPO-R
TPO (R&D) - Parental
O.D. 570
1.8
1.6
1.4
1.2
1
0.8
0.6
0.4
0.2
0
1E-10
1E-9
1E-8
1E-7
Log conc. TPO

FIG. 3A

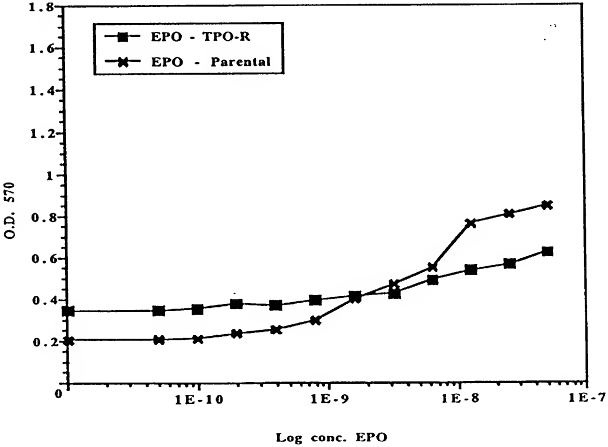

EPO - TPO-R
EPO - Parental
O.D. 570
1.8
1.6
1.4
1.2
1
0.8
0.6
0.4
0.2
0
1E-10
1E-9
1E-8
1E-7
Log conc. EPO

FIG. 3B

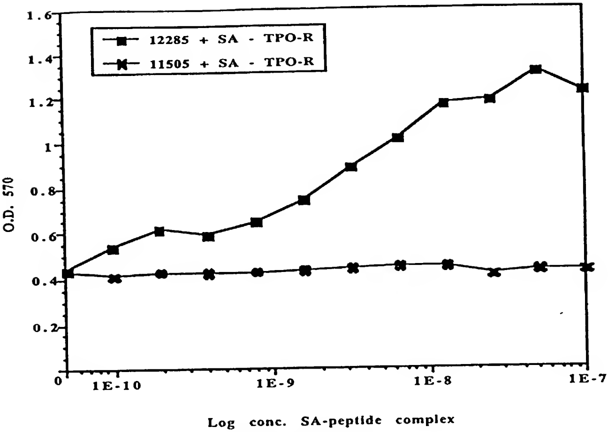

1.6
1.4
1.2
1
0.8
0.6
0.4
0.2
0
O.D. 570
12285 + SA - TPO-R
11505 + SA - TPO-R
1E-10
1E-9
1E-8
1E-7
Log conc. SA-peptide complex

FIG. 3C

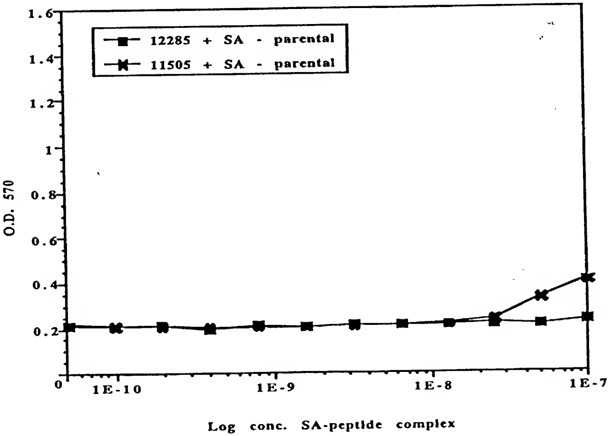

1.6
1.4
1.2
1
0.8
0.6
0.4
0.2
0
O.D. 570
12285 + SA - parental
11505 + SA - parental
1E-10
1E-9
1E-8
1E-7
Log conc. SA-peptide complex

FIG. 3D

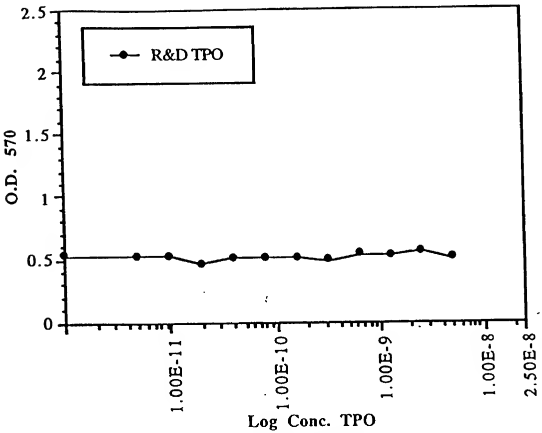

R&D TPO
O.D. 570
2.5
2
1.5
1
0.5
0
1.00E-11
1.00E-10
1.00E-9
1.00E-8
2.50E-8
Log Conc. TPO

FIG. 3E

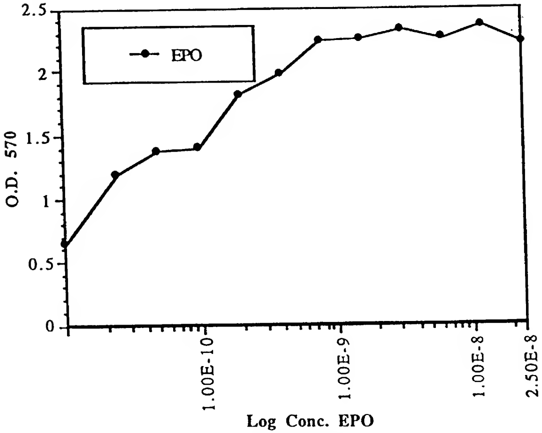

EPO
O.D. 570
2.5
2
1.5
1
0.5
0
1.00E-10
1.00E-9
1.00E-8
2.50E-8
Log Conc. EPO

FIG. 3F

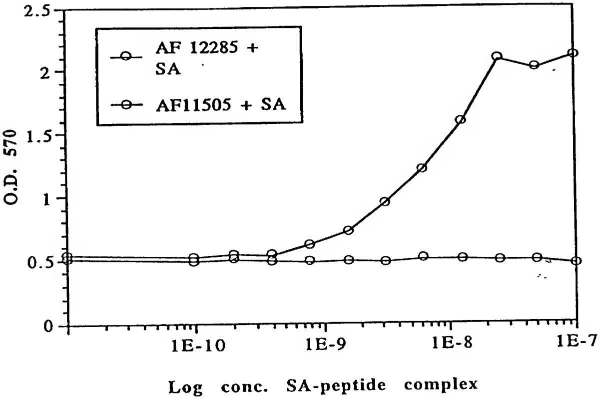

2.5
2
1.5
1
0.5
0
O.D. 570
AF 12285 + SA
AF11505 + SA
1E-10
1E-9
1E-8
1E-7
Log conc. SA-peptide complex

FIG. 3G

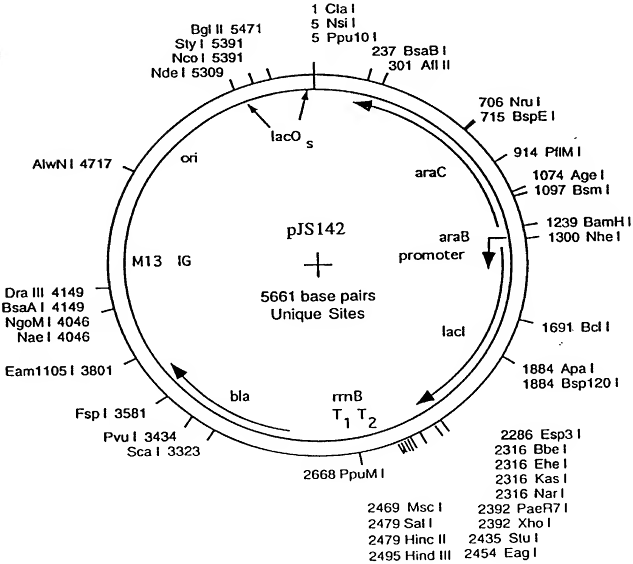
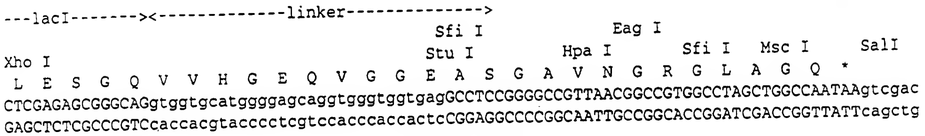

1 Cla I
5 Nsi I
5 Ppu10 I
237 BsaB I
301 Afl II
706 Nru I
715 BspE I
914 PflM I
1074 Age I
1097 Bsm I
1239 BamH I
1300 Nhe I
1691 Bcl I
1884 Apa I
1884 Bsp120 I
2286 Esp3 I
2316 Bbe I
2316 Ehe I
2316 Kas I
2316 Nar I
2392 PaeR7 I
2392 Xho I
2435 Stu I
2454 Eag I
2469 Msc I
2479 Sal I
2479 Hinc II
2495 Hind III
2668 PpuM I
Sca I 3323
Pvu I 3434
Fsp I 3581
Eam1105 I 3801
Nae I 4046
NgoM I 4046
BsaA I 4149
Dra III 4149
AlwN I 4717
Nde I 5309
Nco I 5391
Sty I 5391
Bgl II 5471
lacO s
ori
araC
araB promoter
lacI
rrnB T_1 T_2
bla
M13 IG
pJS142
5661 base pairs
Unique Sites

FIG. 4A

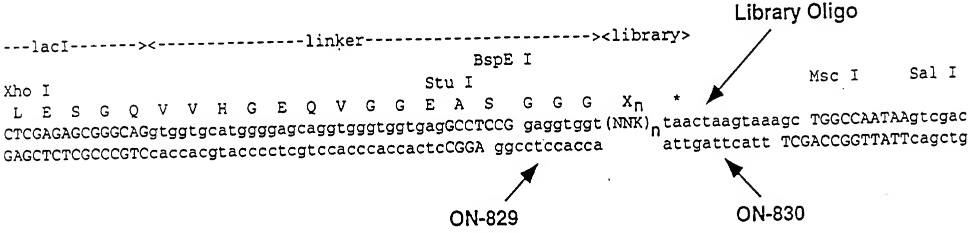
pJS142 Library Vector, cloning sites at 3' end of lacI gene:



```
---lacI------><-------------linker-------------->
                                                Sfi I          Eag I
                                               Stu I      Hpa I      Sfi I  Msc I     SalI
Xho I
 L  E  S  G  Q  V  V  H  G  E  Q  V  G  G  E  A  S  G  A  V  N  G  R  G  L  A  G  Q  *
CTCGAGAGCGGGCAGgtggtgcatggggagcaggtgggtggtgagGCCTCCGGGGCCGTTAACGGCCGTGGCCTAGCTGGCCAATAAgtcgac
GAGCTCTCGCCCGTCcaccacgtacccctcgtccacccaccactcCGGAGGCCCCGGCAATTGCCGGCACCGGATCGACCGGTTATTcagctg
```

FIG. 4B

library Construction after SfiI digestion:



```
                                                                                    Library Oligo
---lacI------><--------------linker----------------------><library>
                                                BspE I                                       Msc I    Sal I
                                            Stu I
Xho I
 L  E  S  G  Q  V  V  H  G  E  Q  V  G  G  E  A  S   G  G  G  Xn    *
CTCGAGAGCGGGCAGgtggtgcatggggagcaggtgggtggtgagGCCTCCG gaggtggt(NNK)n taactaagtaaagc TGGCCAATAAgtcgac
GAGCTCTCGCCCGTCcaccacgtacccctcgtccacccaccactcCGGA ggcctccacca           attgattcatt TCGACCGGTTATTcagctg
                                                       ON-829                   ON-830
```

FIG. 4C

Applicant(s): DOWER et al.
Serial No.: To be assigned
Attorney: Virginia C. Bennett
Telephone: (919) 483-1012
Sheet 10 of 15

pELM3/pELM15 MBP vector cloning sites:

```
---MBP----> <---------------------------linker---------------------------> <----Xa
                    Sac I
 Q   T   N   S   S   S   N   N   N   N   N   N   N   N   N   N   L   G   I   E
CAG ACT AAT TCG AGC TCG AAC AAC AAC AAC AAT AAC AAT AAC AAC AAC CTC GGG ATC GAG
GTC TGA TTA AGC TCG AGC TTG TTG TTG TTG TTA TTG TTA TTG TTG TTG GAG CCC TAG CTC

Xa----> <----------------------------cloning sites---------------------------->
        Age I   Pml I    Sma I  EcoR I  BamH I  Xba I   Sal I   Pst I    Hind III
 G   R   T   G   H   V   A   R   E   F   G   S   S   R   V   D   L   Q   A   S
GGA AGG ACC GGT CAC GTG GCC CGG GAA TTC GGA TCC TCT AGA GTC GAC CTG CAG GCA AGC TT
CCT TCC TGG CCA GTG CAC CGG GCC CTT AAG CCT AGG AGA TCT CAG CTG GAC GTC CGT TCG AA
```

FIG. 5A

pELM3/pELM15 after subcloning of library insert:

```
Xa----> <----linker---> <library>
                                                  Msc I           Sal I
 G   R   T   G   G   G     Xn     *
GGA AGG ACC GGA GGT GGT (NNK)n TAA CTA AGT AAA GCT GGC CAA TAA GTC GAC
CCT TCC TGG CCT CCA CCA (NNM)n ATT GAT TCA TTT CGA CCG GTT ATT CAG CTG
```

FIG. 5B

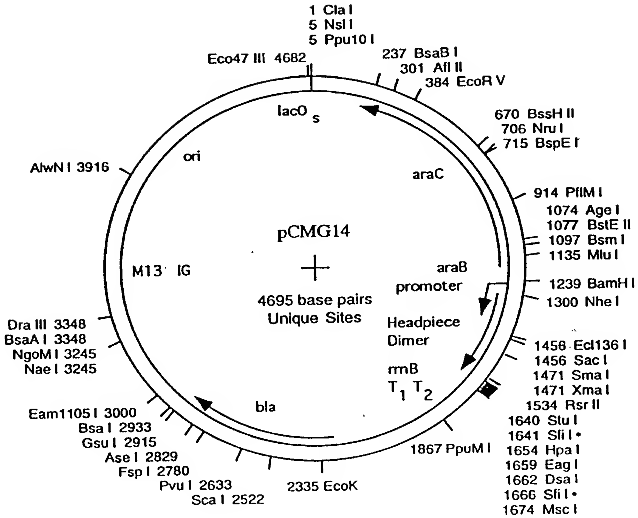

1 Cla I
5 Nsl I
5 Ppu10 I
Eco47 III 4682
237 BsaB I
301 Afl II
384 EcoR V
670 BssH II
706 Nru I
715 BspE I
914 PflM I
1074 Age I
1077 BstE II
1097 Bsm I
1135 Mlu I
1239 BamH I
1300 Nhe I
1458 Ecl136 I
1456 Sac I
1471 Sma I
1471 Xma I
1534 Rsr II
1640 Stu I
1641 Sfi I •
1654 Hpa I
1659 Eag I
1662 Dsa I
1666 Sfi I •
1674 Msc I
1684 Acc I
1684 Sal I
1688 BspM I
1689 Sse8337 I
1696 Sph I
1702 Hind III
1867 PpuM I
2335 EcoK
Sca I 2522
Pvu I 2633
Fsp I 2780
Ase I 2829
Gsu I 2915
Bsa I 2933
Eam1105 I 3000
Nae I 3245
NgoM I 3245
BsaA I 3348
Dra III 3348
AlwN I 3916
lacO s
ori
araC
araB promoter
Headpiece Dimer
rrnB T1 T2
bla
M13 IG
pCMG14
4695 base pairs
Unique Sites

FIG. 6A

pCMG14 Library Vector, cloning sites at 3' end of Headpiece Dimer gene:

```
-----------Headpiece------------><----linker----->
                                           Sfi I            Eag I
                                          Stu I         Hpa I     Sfi I   Msc I    SalI
 E  A  A  M  A  E  L  N  Y  I  P  R  S  Q  E  A  S  G  A  V  N  G  R  G  L  A  G  Q  *
GAAGCGGCGATGGCGGAGCTGAATTACATTCCCcggtcgcaggagGCCTCCGGGGCCGTTAACGGCCGTGGCCTAGCTGGCCAATAAgtcgac
CTTCGCCGCTACCGCCTCGACTTAATGTAAGGGgccagcgtcctcCGGAGGCCCCGGCAATTGCCGGCACCGGATCGACCGGTTATTcagctg
```

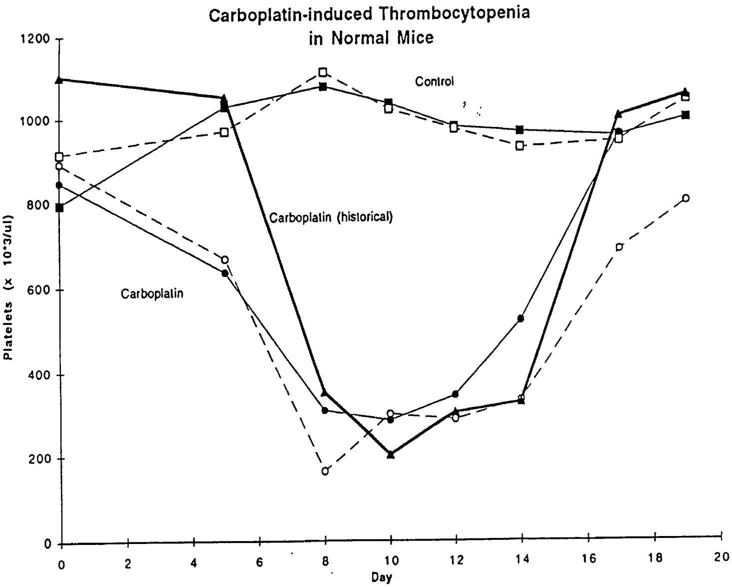
FIG. 6B

Library Construction after SfiI digestion:

```
                                                                           ON-1679
-----------Headpiece------------><-------linker------------><library>
                                                 BspE I
Xho I                                     Stu I                                      Msc I     Sal I
 E  A  A  M  A  E  L  N  Y  I  P  R  S  Q  E  A  S   G  G  G  X12  *
GAAGCGGCGATGGCGGAGCTGAATTACATTCCCcggtcgcaggagGCCTCCG gaggtggt(NNK)12 taactaagtaaagc TGGCCAATAAgtcgac
CTTCGCCGCTACCGCCTCGACTTAATGTAAGGGgccagcgtcctcCGGA ggcctccacca        attgattcatt TCGACCGGTTATTcagctg

                                                ON-829                     ON-830
```

FIG. 6C



Carboplatin-induced Thrombocytopenia in Normal Mice

BALB/c mice were treated wilh carboplatin (125 mg/kg) on Day 0.

Dashed lines represent female mice.

Solid lines represent male mice.

Bold line represents data published by Ulich, T. R., et al. Blood 86(3):971-976. 1995.

FIG. 7

Carboplatin-induced Thrombocytopenia in Normal Mice:
Single-dose Titration (mg/kg)

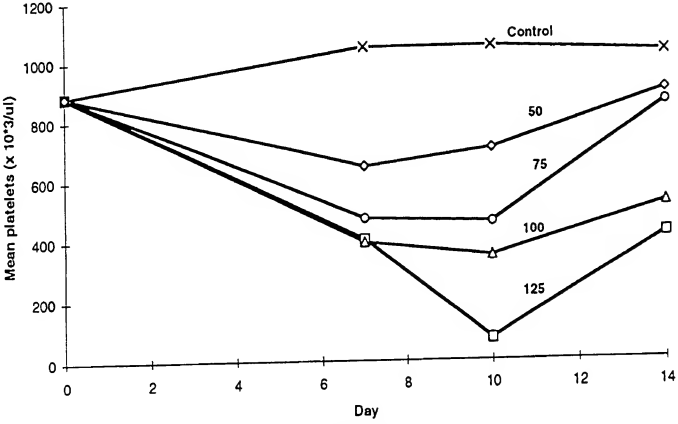
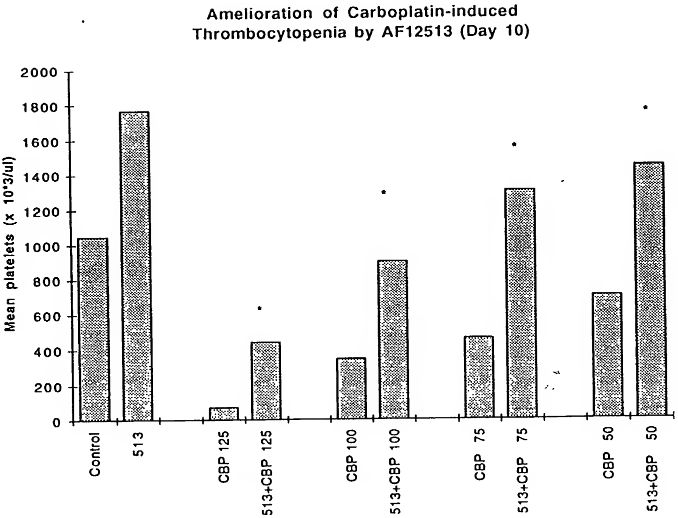


FIG. 8



Carboplatin (CBP; 125-50 mg/kg, ip) was given on Day 0.
AF12513 (513; 1 mg/kg, ip) was given on Days 1-9.

* p= <0.05

FIG. 9